### (19) World Intellectual Property Organization International Bureau



## . DOOD BILLEON AND BILLOON AND A CONTRACTOR OF THE CONTRACTOR OF T

(43) International Publication Date 19 May 2005 (19.05.2005)

#### PCT

# (10) International Publication Number WO 2005/044086 A2

(51) International Patent Classification7:

**A61B** 

(21) International Application Number:

PCT/US2004/035929

(22) International Filing Date: 29 October 2004 (29.10.2004)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/515,735 30 O

30 October 2003 (30.10.2003)

- (71) Applicant (for all designated States except US): TUFTS-NEW ENGLAND MEDICAL CENTER [US/US]; 750 Washington Street, Box 394, Boston, MA 02111 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): BIANCHI, Diana, W. [US/US]; 4 Perry Street, Brookline, MA 02445 (US). LARRABEE, Paige, B. [US/US]; 19 Cherry Street, Malden, MA 02148 (US). LeSHANE, Erik, S. [US/US]; 131 Madbury Road, Durham, NH 03824 (US).
- (74) Agent: JARRELL, Brenda, Herschbach; Choate, Hall & Stewart, Exchange Place, 53 State Street, Boston, MA 02109 (US).

- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

#### Published:

 without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: PRENATAL DIAGNOSIS USING CELL-FREE FETAL DNA IN AMNIOTIC FLUID

(57) Abstract: The present invention relates to improved methods of prenatal diagnosis, screening, monitoring and/or testing. The inventive methods include the analysis by array-based hybridization of cell-free fetal DNA isolated from amniotic fluid. In addition to allowing the prenatal diagnosis of a variety of diseases and conditions, and the assessment of fetal characteristics such as fetal sex and chromosomal abnormalities, the new inventive methods provide substantially more information about the fetal genome in less time than it takes to perform a conventional metaphase karyotype analysis. In particular, the enhanced molecular karyotype methods provided by the present invention allow the detection of chromosomal aberrations that are not often detected prenatally such as microdeletions, microduplications and subtelomeric rearrangements.

